

CLAIMS

What is claimed is:

1. A compound comprising a peptide chain approximately 10 to 40 amino acids in length that binds to G-CSFR and contains a sequence of amino acids of formula (I)

5 (I) $CX_1X_2X_3X_4X_5X_6X_7X_8C$ (SEQ ID NO: 1)

wherein each amino acid is indicated by standard one-letter abbreviation, and wherein X_1 is A, N, S, F, D, G, L, T, E, V, P, Q, H, M or K; X_2 is M, G, R, H, D, I, V, A, S, E, N, F, Y, P, C, W or T; X_3 is E, V, W, F, M, A, N, S, L, T, Y, G or P; X_4 is V, I, G, Q, W, M, T, Y, L, P, D, C, E or A; X_5 is M, E, W, L, P, N, I, T, V, F, Y, Q, S, R, W, G, H or D; X_6 is
10 H, A, W, Y, V, F, Q, M, N, E, S, D, P or G; X_7 is M, F, Y, V, N, L, H, D, S, W, G, Q, C or T; and X_8 is C, Y, R, I, K, W, L, E, M, H, A, T, F, D, P, G or Q.

2. The compound of claim 1, wherein X_1 is D or P.

15 3. The compound of claim 1, wherein X_2 is D or P.

4. The compound of claim 1, wherein X_3 is E or W.

20 5. The compound of claim 1, wherein X_4 is V, I or Y.

6. The compound of claim 1, wherein X_5 is M or L.

7. The compound of claim 1, wherein X_6 is W, Y or F.

25 8. The compound of claim 1, wherein X_7 is M, Y or D.

9. The compound of claim 1, wherein X_8 is C or M.

10. The compound of claim 1, wherein the sequence of amino acids is selected from the group consisting of:

- 5 CAGEVMHMCC (SEQ ID NO: 8);
CNREIEAMCC (SEQ ID NO: 9);
CADEVMHFCC (SEQ ID NO: 10);
CNREIMWMCC (SEQ ID NO: 11);
CSHEVWWYCC (SEQ ID NO: 12);
CSREVLYYCC (SEQ ID NO: 13);
CFIEGPWVCC (SEQ ID NO: 14);
10 CFVEGNWYCC (SEQ ID NO: 15);
CAAHEVMVNCC (SEQ ID NO: 16);
CSDEVIFYCC (SEQ ID NO: 17);
CDREIMWFCC (SEQ ID NO: 18);
CAHEVMWMCC (SEQ ID NO: 19);
15 CGSEVTFMCC (SEQ ID NO: 20);
CLEEIMWLCC (SEQ ID NO: 21);
CAREVLAMCC (SEQ ID NO: 22);
CSVEVMQMCC (SEQ ID NO: 23);
CTNVQLMHYC (SEQ ID NO: 24);
20 CDVWQLFDRC (SEQ ID NO: 25);
CSFVQLNSIC (SEQ ID NO: 26);
CDYWQWFDKC (SEQ ID NO: 27);
CESFWVELWC (SEQ ID NO: 28);
CVPWMFYDLC (SEQ ID NO: 29);
25 CDPWMFYDLC (SEQ ID NO: 30);
CDPWVLFDEC (SEQ ID NO: 31);
CDHWTYFDMC (SEQ ID NO: 32);
CVVWTLYDKC (SEQ ID NO: 33);

CPDWYQSYMC (SEQ ID NO: 34);
CPDWYSYYMC (SEQ ID NO: 35);
CPEWYTDVMC (SEQ ID NO: 36);
CPDWYLDYMC (SEQ ID NO: 37);
5 CPEWYLDYMC (SEQ ID NO: 38);
CPDWYLPYMC (SEQ ID NO: 39);
CPEWYLPYMC (SEQ ID NO: 40);
CQDWWVELWC (SEQ ID NO: 41);
CPDWYLPWMC (SEQ ID NO: 42);
10 CACMLRVVHC (SEQ ID NO: 43);
CQRAGYMLAC (SEQ ID NO: 44);
CHANPVWGEC (SEQ ID NO: 45);
CFWSDWGQTC (SEQ ID NO: 46);
CPHWTSYYMC (SEQ ID NO: 47);
15 CETLCGACFC (SEQ ID NO: 48);
CATTINDTLC (SEQ ID NO: 49);
CLNYPHPVFC (SEQ ID NO: 50);
CMDGEMAVDC (SEQ ID NO: 51);
CNMGWMSWPC (SEQ ID NO: 52);
20 CETYADWLGC (SEQ ID NO: 53);
CDPWMFFDMC (SEQ ID NO: 54);
CDPWIWYDLC (SEQ ID NO: 55);
CDPWIMYDRC (SEQ ID NO: 56);
CDPWVFFDIC (SEQ ID NO: 57);
25 CDPWTYYDLC (SEQ ID NO: 58);
CDPWIFYDRC (SEQ ID NO: 59);
CDPWLFYDLC (SEQ ID NO: 60);
CDPWVWYDLC (SEQ ID NO: 61);

CDPWIFFDRC (SEQ ID NO: 62);
CDPWMFFDQC (SEQ ID NO: 63);
CDPWLWYDRC (SEQ ID NO: 64);
CDVWVWYDQC (SEQ ID NO: 65);
5 CDPWIYYDLC (SEQ ID NO: 66);
CVPWTLFDLC (SEQ ID NO: 67);
CPAWYLEYMC (SEQ ID NO: 68);
CPDWYLEYMC (SEQ ID NO: 69);
CKYWQWFDKC (SEQ ID NO: 70); and
10 CDHWMWYDKC (SEQ ID NO: 71).

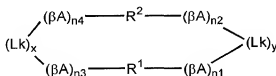
11. The compound of claim 10, wherein the sequence of amino acids is selected from the group consisting of:

GCNREIAMCCG (SEQ ID NO: 72);
15 GCPEWYTDVMCG (SEQ ID NO: 73);
NWYCMDGEMAVDCEAT (SEQ ID NO: 74);
WQSCNMGWMSWPCYFV (SEQ ID NO: 75);
HELCETYADWLGCVEW (SEQ ID NO: 76);
PCDPWMFFDMCERW (SEQ ID NO: 77);
20 LRGCDPWIWYDLCPAV (SEQ ID NO: 78);
GYLCDPWIXYDRCLGF (SEQ ID NO: 79);
RFACDPWVFFDICGYW (SEQ ID NO: 80);
GYWCDPWITYDLCLTA (SEQ ID NO: 81);
MWTCDPWIFYDRCLFN (SEQ ID NO: 82);
25 GSSCDPWLFYDLCLLD (SEQ ID NO: 83);
GGGCDPWVWYDLCWCD (SEQ ID NO: 84);
YTSCDPWIFFDRMSV (SEQ ID NO: 85);
DPYCDPWMFFDQCAYL (SEQ ID NO: 86);

REFCDPWLWYDRCL (SEQ ID NO: 87);
NTGCDVWVWYDQCFAM (SEQ ID NO: 88);
LVFCDPWIIYDLCMDT (SEQ ID NO: 89);
GCSFVQLNSICG (SEQ ID NO: 90);
5 GCPAWLEYMCG (SEQ ID NO: 91);
GCPDWLEYMCG (SEQ ID NO: 92);
GCKYWQWFDKCG (SEQ ID NO: 93); and
GCDHWMWYDKCG (SEQ ID NO: 94).

- 10 12. The compound of claim 1, comprising a dimer having the structure of formula (VIII)

(VIII)



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wherein R^1 and R^2 are independently selected from the sequences of amino acids of formula (I); βA is a β -alanine residue; n_1, n_2, n_3, n_4, x and y are independently zero or one with the proviso that the sum of x and y is either one or two; and Lk is a terminal linking moiety selected from the group consisting of a disulfide bond, a carbonyl moiety, 20 a C_{1-12} linking moiety optionally terminated with one or two $-\text{NH}-$ linkages and optionally substituted at one or more available carbon atoms with a lower alkyl substituent, a lysine residue or a lysine amide.

13. The compound of claim 1, containing a disulfide bond.

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14. The compound of claim 1, wherein the N-terminus of the peptide is coupled to a polyethylene glycol molecule.

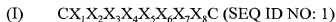
15. The compound of claim 1, wherein the N-terminus of the peptide is acetylated.

16. The compound of claim 1, wherein the C-terminus of the peptide is amidated.

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17. A pharmaceutical composition comprising a therapeutically effective amount of the compound of claim 1 in combination with a pharmaceutically acceptable carrier.

18. A method for treating a patient who would benefit from administration of a
10 G-CSF modulator, comprising administering to the patient a therapeutically effective amount of a compound comprising a peptide chain approximately 10-40 amino acids in length that binds to G-CSFR and contains a sequence of amino acids having the structural formula (I)



15 wherein each amino acid is indicated by standard one-letter abbreviation, and wherein X_1 is A, N, S, F, D, G, L, T, E, V, P, Q, H, M or K; X_2 is M, G, R, H, D, I, V, A, S, E, N, F, Y, P, C, W or T; X_3 is E, V, W, F, M, A, N, S, L, T, Y, G or P; X_4 is V, I, G, Q, W, M, T, Y, L, P, D, C, E or A; X_5 is M, E, W, L, P, N, I, T, V, F, Y, Q, S, R, W, G, H or D; X_6 is H, A, W, Y, V, F, Q, M, N, E, S, D, P or G; X_7 is M, F, Y, V, N, L, H, D, S, W, G, Q, C
20 or T; and X_8 is C, Y, R, I, K, W, L, E, M, H, A, T, F, D, P, G or Q.

19. The method of claim 18, wherein the G-CSF modulator is an agonist for the G-CSFR.

25 20. The method of claim 19, wherein the patient suffers from a depressed neutrophil count.

21. The method of claim 20, wherein the depressed neutrophil count is caused by a condition selected from the group consisting of chemotherapy-induced neutropenia, AIDS-induced neutropenia and community-acquired pneumonia-induced neutropenia.

5 22. The method of claim 18, wherein the G-CSF modulator is an antagonist for the G-CSFR.

23. A compound comprising a peptide chain approximately 9 to 40 amino acids in length that binds to G-CSFR and contains a sequence of amino acids of formula (II)

10 (II) $X_1^1 X_2^1 X_3^1 \text{SGWVWX}_4^1$ (SEQ ID NO: 2)

wherein each amino acid is indicated by the standard one-letter abbreviation, and wherein X_1^1 is S, Q, R, L or Y; X_2^1 is N, S, T, A or D; X_3^1 is E, D or N; and X_4^1 is L, V, T, P or H.

15 24. The compound of claim 23, wherein X_1^1 is S or Q.

25. The compound of claim 23, wherein X_2^1 is S.

26. The compound of claim 23, wherein X_3^1 is N.

20 27. The compound of claim 23, wherein X_4^1 is V.

28. The compound of claim 23, wherein the sequence of amino acids is selected from the group consisting of:

25 SNESGWVWL (SEQ ID NO: 95);
 QNSNGWVWV (SEQ ID NO: 96);
 RTESGWVWT (SEQ ID NO: 97);
 RANSWVWV (SEQ ID NO: 98);
 YDNSGWVWH (SEQ ID NO: 99); and

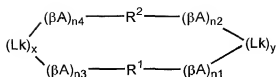
LSDSGWVWVP (SEQ ID NO: 100).

29. The compound of claim 28, wherein the sequence of amino acids is selected from the group consisting of:

- 5 EQNSGWWVVGGGGC (SEQ ID NO: 101);
CEQNSGWWV (SEQ ID NO: 102);
EQNSGWWVVGGGGCKKK (SEQ ID NO: 103);
EQNSGWWVVGKKKC (SEQ ID NO: 104);
EQNSGWWVVGKKK (SEQ ID NO: 105);
10 KKKEQNSGWWV (SEQ ID NO: 106);
EQNSGWWVVGKKKSKKK (SEQ ID NO: 107);
EQNSGWWVVGCKKK (SEQ ID NO: 108);
EQNSGWWVVGGGGGCKKK (SEQ ID NO: 109);
SNESGWWLP (SEQ ID NO: 110);
15 EQNSGWWV (SEQ ID NO: 111);
SRTEGWWT (SEQ ID NO: 112);
QRANSWWV (SEQ ID NO: 113);
DYDNSGWWH (SEQ ID NO: 114).
EQNSGWWVVGKKK (SEQ ID NO: 115);
20 EQNSGWWVVGGGGSKKK (SEQ ID NO: 116);
EQNSGWWVVGGGGS (SEQ ID NO: 117);
EQNSGWWVVGGGGSEQNSGWWVVGGGGS (SEQ ID NO: 118);
RYQSFELSDSGWVWVPVARH (SEQ ID NO: 119); and
EQNSGWWVVGGGGCKKK (SEQ ID NO: 492)
- 25

30. The compound of claim 23, comprising a dimer having the structure of formula (VIII)

(VIII)



- 5 wherein R^1 and R^2 are independently selected from the sequences of amino acids of formula (II); βA is a β -alanine residue; $n1$, $n2$, $n3$, $n4$, x and y are independently zero or one with the proviso that the sum of x and y is either one or two; and Lk is a terminal linking moiety selected from the group consisting of a disulfide bond, a carbonyl moiety, a C_{1-12} linking moiety optionally terminated with one or two $-\text{NH}-$ linkages and optionally substituted at one or more available carbon atoms with a lower alkyl substituent, a lysine residue or a lysine amide.

31. The compound of claim 30, wherein the dimer is:

$\text{NH}_2\text{-EQSNSGWVWVGGGGC-CONH}_2$ (SEQ ID NO: 101)

$\text{NH}_2\text{-EQSNSGWVWVGGGGC-CONH}_2$ (SEQ ID NO: 101);

32. The compound of claim 23, containing a disulfide bond.

- 20 33. The compound of claim 23, wherein the N-terminus of the peptide is coupled to a polyethylene glycol molecule.

34. The compound of claim 23, wherein the N-terminus of the peptide is acetylated.

25 35. The compound of claim 23, wherein the C-terminus of the peptide is amidated.

36. A pharmaceutical composition comprising a therapeutically effective amount of the compound of claim 23 in combination with a pharmaceutically acceptable carrier.

37. A method for treating a patient who would benefit from administration of a
5 G-CSF modulator, comprising administering to the patient a therapeutically effective amount of a compound comprising a peptide chain approximately 9 to 40 amino acids that binds to G-CSF and contains a sequence of amino acids having the structural formula (II)

(II) $X_1^I X_2^I X_3^I SGWVWX_4^I$ (SEQ ID NO: 2)

10 wherein each amino acid is indicated by the standard one-letter abbreviation, and wherein X_1^I is S, Q, R, L or Y; X_2^I is N, S, T, A or D; X_3^I is E, D or N; and X_4^I is L, V, T, P or H.

38. The method of claim 37, wherein the G-CSF modulator is an agonist for the G-CSFR.

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39. The method of claim 38, wherein the patient suffers from a depressed neutrophil count.

40. The method of claim 39, wherein the depressed neutrophil count is caused by
20 a condition selected from the group consisting of chemotherapy-induced neutropenia, AIDS-induced neutropenia and community-acquired pneumonia-induced neutropenia.

41. The method of claim 37, wherein the G-CSF modulator is an antagonist for the G-CSFR.

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42. A compound comprising a peptide chain approximately 6 to 40 amino acids in length that binds to G-CSFR and contains a sequence of amino acids of formula (III)

(III) $ERX^{II}_1 X_2^{II} X_3^{II} C$ (SEQ ID NO: 3)

wherein each amino acid is indicated by standard one-letter abbreviation, and wherein X^{II}_1 is D, L, S, G, E, A, K or Y; X^{II}_2 is W, Y, F, L or V; and X^{II}_3 is F, G, M or L.

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43. The compound of claim 42, wherein X^{II}_1 is D or L.

44. The compound of claim 42, wherein X^{II}_2 is W.

45. The compound of claim 42, wherein X^{II}_3 is F.

10

46. The compound of claim 42, wherein the sequence of amino acids is selected from the group consisting of:

ERDWFC (SEQ ID NO: 120);

ERDWGC (SEQ ID NO: 121);

ERLWFC (SEQ ID NO: 122);

15

ERSYFC (SEQ ID NO: 123);

ERGWFC (SEQ ID NO: 124);

EREWFC (SEQ ID NO: 125);

ERAWFC (SEQ ID NO: 126);

ERLYFC (SEQ ID NO: 127);

20

ERYFMC (SEQ ID NO: 128);

ERLFLC (SEQ ID NO: 129);

ERALMC (SEQ ID NO: 130);

ERDVMC (SEQ ID NO: 131); and

ERKWFC (SEQ ID NO: 132).

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47. The compound of claim 46, wherein the sequence of amino acids is selected from the group consisting of:

ETWGERDWFC (SEQ ID NO: 133);

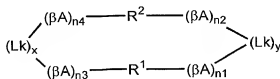
ETWGERDWGC (SEQ ID NO: 134);
STAERLWFCG (SEQ ID NO: 135);
YETAERSYFC (SEQ ID NO: 136);
ADNAERGWFC (SEQ ID NO: 137);
5 QNSEREWFC (SEQ ID NO: 138);
STSERAWFCG (SEQ ID NO: 139);
ASWSEGWFC (SEQ ID NO: 140);
ELSSEREWFC (SEQ ID NO: 141);
DMQGERGWFC (SEQ ID NO: 142);
10 SSSERAWFCG (SEQ ID NO: 143);
GNMRERLYFC (SEQ ID NO: 144);
QPNRERYFMC (SEQ ID NO: 145);
SVTRERLFLC (SEQ ID NO: 146);
IPLSERALMCSSWNC (SEQ ID NO: 147);
15 WARSERDVMCLSYVC (SEQ ID NO: 148);
QNSEREWFCG (SEQ ID NO: 149);
QNSEREWFCGGGS (SEQ ID NO: 150);
NLEEALAQERLWFCRSGNC (SEQ ID NO: 151); and
NLESYEMEERKWFCCKMFSC (SEQ ID NO: 152).

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48. The compound of claim 42, comprising a dimer having the structure of
formula (VIII)

(VIII)

25



wherein R¹ and R² are independently selected from the sequences of amino acids of
formula (III); βA is a β-alanine residue; n1, n2, n3, n4, x and y are independently zero or

one with the proviso that the sum of x and y is either one or two; and Lk is a terminal linking moiety selected from the group consisting of a disulfide bond, a carbonyl moiety, a C₁₋₁₂ linking moiety optionally terminated with one or two -NH- linkages and optionally substituted at one or more available carbon atoms with a lower alkyl substituent, a lysine residue or a lysine amide.

49. The compound of claim 42, containing a disulfide bond.

50. The compound of claim 49, selected from the group consisting of:

NH₂-STAERLWFCG-CONH₂ (SEQ ID NO: 135)

NH₂-STAERLWFCG-CONH₂ (SEQ ID NO: 135);

NH₂-QNSSEREWFC-CONH₂ (SEQ ID NO: 138)

NH₂-QNSSEREWFC-CONH₂ (SEQ ID NO: 138); and

NH₂-QNSSEREWFCG-CONH₂ (SEQ ID NO: 149)

NH₂-QNSSEREWFCG-CONH₂ (SEQ ID NO: 149).

51. The compound of claim 42, wherein the N-terminus of the peptide is coupled to a polyethylene glycol molecule.

52. The compound of claim 42, wherein the N-terminus of the peptide is acetylated.

53. The compound of claim 42, wherein the C-terminus of the peptide is amidated.

5 G-CSF modulator, comprising administering to the patient a therapeutically effective amount of a compound comprising a peptide chain approximately 6 to 40 amino acids that binds to G-CSFR and contains a sequence of amino acids having the structural formula (III)

10 wherein each amino acid is indicated by standard one-letter abbreviation, and wherein X^{II} ,
is D, L, S, G, E, A, K or Y; X^{II}_2 is W, Y, F, L or V; and X^{II}_3 is F, G, M or L.

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57. The method of claim 56, wherein the patient suffers from a depressed neutrophil count.

59. The method of claim 55, wherein the G-CSF modulator is an antagonist for the G-CSFR.

60. A compound comprising a peptide chain approximately 9 to 40 amino acids in length that binds to G-CSFR and contains a sequence of amino acids of formula (IV)

(IV) $X^{III_1}MVYX^{III_2}X^{III_3}PX^{III_4}W$ (SEQ ID NO: 4)

wherein each amino acid is indicated by standard one-letter abbreviation, and wherein X^{III}_1 is D or E; X^{III}_2 is A or T; X^{III}_3 is Y or V; and X^{III}_4 is P or Y.

61. The compound of claim 60, wherein the sequence of amino acids is selected
5 from the group consisting of:

DMVYAYPPW (SEQ ID NO: 153); and

EMVYTVPYW (SEQ ID NO: 154).

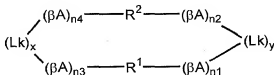
62. The compound of claim 61, wherein the sequence of amino acids is selected
10 from the group consisting of:

DMVYAYPPWS (SEQ ID NO: 155); and

DEMVYTVPYW (SEQ ID NO: 156).

63. The compound of claim 60, comprising a dimer having the structure of
15 formula (VIII)

(VIII)



- 20 wherein R^1 and R^2 are independently selected from the sequences of amino acids of formula (IV); βA is a β -alanine residue; n_1 , n_2 , n_3 , n_4 , x and y are independently zero or one with the proviso that the sum of x and y is either one or two; and Lk is a terminal linking moiety selected from the group consisting of a disulfide bond, a carbonyl moiety, a C_{1-12} linking moiety optionally terminated with one or two $-NH-$ linkages and optionally
25 substituted at one or more available carbon atoms with a lower alkyl substituent, a lysine residue or a lysine amide.

64. The compound of claim 60, containing a disulfide bond.

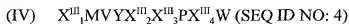
65. The compound of claim 60, wherein the N-terminus of the peptide is coupled to a polyethylene glycol molecule.

66. The compound of claim 60, wherein the N-terminus of the peptide is
5 acetylated.

67. The compound of claim 60, wherein the C-terminus of the peptide is amidated.

10 68. A pharmaceutical composition comprising a therapeutically effective amount of the compound of claim 60 in combination with a pharmaceutically acceptable carrier.

69. A method for treating a patient who would benefit from administration of a G-CSF modulator, comprising administering to the patient a therapeutically effective
15 amount of a compound comprising a peptide chain approximately 9 to 40 amino acids that binds to G-CSFR and contains a sequence of amino acids having the structural formula (IV)



wherein each amino acid is indicated by standard one-letter abbreviation, and wherein
20 X^{III}_1 is D or E; X^{III}_2 is A or T; X^{III}_3 is Y or V; and X^{III}_4 is P or Y.

70. The method of claim 69, wherein the G-CSF modulator is an agonist for the G-CSFR.

25 71. The method of claim 70, wherein the patient suffers from a depressed neutrophil count.

72. The method of claim 71, wherein the depressed neutrophil count is caused by a condition selected from the group consisting of chemotherapy-induced neutropenia, AIDS-induced neutropenia and community-acquired pneumonia-induced neutropenia.

5 73. The method of claim 69, wherein the G-CSF modulator is an antagonist for the G-CSFR.

Solo 74. A compound comprising a peptide chain approximately 12 to 40 amino acids in length that binds to G-CSFR and contains a sequence of amino acids of formula (V)
10 (V) $CX^{IV}_1X^{IV}_2X^{IV}_3X^{IV}_4X^{IV}_5X^{IV}_6X^{IV}_7X^{IV}_8X^{IV}_9X^{IV}_{10}C$ (SEQ ID NO: 5)
wherein each amino acid is indicated by standard one-letter abbreviation, and wherein
 X^{IV}_1 is E, G, P, N, R, T, W, S, L, H, A, Q or Y; X^{IV}_2 is S, T, E, A, D, G, W, P, L, N, V, Y, R or M; X^{IV}_3 is R, Y, V, Q, E, T, L, P, S, K, M, A or W; X^{IV}_4 is L, M, G, F, W, R, S, V, P, A, D, C or T; X^{IV}_5 is V, T, A, R, S, L, W, C, I, E, P, H, F, D or Q; X^{IV}_6 is E, Y, G, T, Q,
15 M, S, N, A or P; X^{IV}_7 is C, V, D, G, L, W, E, V, I, S, M or A; X^{IV}_8 is S, Y, A, W, P, V, L, Q, G, K, F, I, E or D; X^{IV}_9 is R, W, M, D, H, V, G, A, Q, L, S, E or Y; X^{IV}_{10} is M, L, I, S, V, P, W, F, T, Y, R, or Q.

20 75. The compound of claim 74, wherein X^{IV}_1 is E.

76. The compound of claim 74, wherein X^{IV}_2 is S or A.

77. The compound of claim 74, wherein X^{IV}_3 is R.

25 78. The compound of claim 74, wherein X^{IV}_4 is L.

79. The compound of claim 74, wherein X^{IV}_5 is V or S.

80. The compound of claim 74, wherein X^{IV}_6 is E.

81. The compound of claim 74, wherein X^{IV}_7 is C.

5 82. The compound of claim 74, wherein X^{IV}_8 is S.

83. The compound of claim 74, wherein X^{IV}_9 is R.

84. The compound of claim 74, wherein X^{IV}_{10} is L.

10

85. The compound of claim 74, wherein the sequence of amino acids is selected from the group consisting of:

CESRLVECSRMC (SEQ ID NO: 157);

CETYMTYVYWLC (SEQ ID NO: 158);

15

CGERLAECARLC (SEQ ID NO: 159);

CESRLRECSMLC (SEQ ID NO: 160);

CEARLSECSRIC (SEQ ID NO: 161);

CPARLLECSRMC (SEQ ID NO: 162);

CESVGVDWWSC (SEQ ID NO: 163);

20

CEDRLVEGPWVC (SEQ ID NO: 164);

CNDQFRTCVDVC (SEQ ID NO: 165);

CRGEWWELYHPC (SEQ ID NO: 166);

CEDTRTGAWWSC (SEQ ID NO: 167);

CTWLSSGELVWC (SEQ ID NO: 168);

25

CWPPVCEVSGIC (SEQ ID NO: 169);

CSLSPIQLQHLC (SEQ ID NO: 170);

CLARLEECSRFC (SEQ ID NO: 171);

CHNSSPMVGVTC (SEQ ID NO: 172);

CHVSPVQIKALC (SEQ ID NO: 173);
CAAPATSWFQYC (SEQ ID NO: 174);
CASKLHECSLRC (SEQ ID NO: 175);
CEPMDSNGIVQC (SEQ ID NO: 176);
5 CQYASAADEQRC (SEQ ID NO: 177);
CEYWDEPSLSWC (SEQ ID NO: 178);
CERECFQMLERC (SEQ ID NO: 179);
CGMSTDELDEIC (SEQ ID NO: 180);
CYVSPSTGLYSC (SEQ ID NO: 181);
10 CEARLVECSRLC (SEQ ID NO: 182);
CESRLSECSRMC (SEQ ID NO: 183);
CELKLQECARRC (SEQ ID NO: 184);
CELKLQEAARRC (SEQ ID NO: 185); and
CLERLEECSRFC (SEQ ID NO: 186).

15 86. The compound of claim 85, wherein the sequence of amino acid is selected from the group consisting of:

GGCESRLVECSRMC (SEQ ID NO: 187);
GGCETYMTYVYWLC (SEQ ID NO: 188);
20 EWLCEVGVGDWWSC (SEQ ID NO: 189);
YHPCEDRLVEGPWWCCRS (SEQ ID NO: 190);
WLLCNDQFRTCDVCDNV (SEQ ID NO: 191);
IAECRGEWWEYHPCLAA (SEQ ID NO: 192);
TWYCEDTRTGVAWSCLEL (SEQ ID NO: 193);
25 QLDCTWLSSGELVWCSDW (SEQ ID NO: 194);
QFDCTWLSSGELVWCSDW (SEQ ID NO: 195);
CWPPVCEVSGICS (SEQ ID NO: 196);
CGCSLSPIQLQHLC (SEQ ID NO: 197);

CGCHVSPVQIKALC (SEQ ID NO: 198);
GCHVSPVQIKALC (SEQ ID NO: 199);
GTSCAAPATSWFQYCVLP (SEQ ID NO: 200);
RMDCASKLHECSLRCA YA (SEQ ID NO: 201);
5 GVVCEPMDSNGIVQCSMR (SEQ ID NO: 202);
IDVCQYASAADEORCLRI (SEQ ID NO: 203);
NVLCEYWDEPSLSWCLSS (SEQ ID NO: 204);
CQCERECFQMLERC (SEQ ID NO: 205);
FCSCGMSTDELDEICAHW (SEQ ID NO: 206);
10 EEVCYVSPSTGLYSCYDQ (SEQ ID NO: 207);
LLDICELKLQECARRCN (SEQ ID NO: 208);
GGGLLDICELKLQECARRCN (SEQ ID NO: 209);
GRTGGGLLDICELKLQECARRCN (SEQ ID NO: 210);
LGIEGRTGGGLLDICELKLQECARRCN (SEQ ID NO: 211);
15 LLDICELKLQEAARRCN (SEQ ID NO: 212); and
KLLDICELKLQEAARRCN (SEQ ID NO: 213).

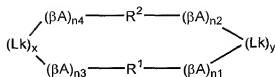
87. The compound of claim 86, wherein the sequence of amino acids is selected from the group consisting of:

20 LLDICELKLQECARRCN (SEQ ID NO: 208);
GGGLLDICELKLQECARRCN (SEQ ID NO: 209);
GRTGGGLLDICELKLQECARRCN (SEQ ID NO: 210);
LGIEGRTGGGLLDICELKLQECARRCN (SEQ ID NO: 211);
LLDICELKLQEAARRCN (SEQ ID NO: 212); and
25 KLLDICELKLQEAARRCN (SEQ ID NO: 213).

88. The compound of claim 74, comprising a dimer having the structure of formula (VIII)

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(VIII)



- 5 wherein R^1 and R^2 are independently selected from the sequences of amino acids of formula (V); βA is a β -alanine residue; $n1$, $n2$, $n3$, $n4$, x and y are independently zero or one with the proviso that the sum of x and y is either one or two; and Lk is a terminal linking moiety selected from the group consisting of a disulfide bond, a carbonyl moiety, a C_{1-12} linking moiety optionally terminated with one or two $-NH-$ linkages and optionally substituted at one or more available carbon atoms with a lower alkyl substituent, a lysine residue or a lysine amide.

89. The compound of claim 74, containing a disulfide bond.

- 15 90. The compound of claim 89, having the structure:
 $NH_3^+ \text{---} LL \text{---} DICELKLQECARRCN \text{---} COO^-$ (SEQ ID NO: 208)
 $NH_3^+ \text{---} LLDICELKLQECARRCN \text{---} COO^-$ (SEQ ID NO: 208).

- 20 91. The compound of claim 74, wherein the N-terminus of the peptide is coupled to a polyethylene glycol molecule.

92. The compound of claim 74, wherein the N-terminus of the peptide is acetylated.

25

93. The compound of claim 74, wherein the C-terminus of the peptide is amidated.

94. A pharmaceutical composition comprising a therapeutically effective amount of the compound of claim 74 in combination with a pharmaceutically acceptable carrier.

95. A method for treating a patient who would benefit from administration of a
- 5 G-CSF modulator, comprising administering to the patient a therapeutically effective amount of a compound comprising a peptide chain approximately 12 to 40 amino acids that binds to G-CSFR and contains a sequence of amino acids having the structural formula (V)
- (V) $CX^{IV}_1X^{IV}_2X^{IV}_3X^{IV}_4X^{IV}_5X^{IV}_6X^{IV}_7X^{IV}_8X^{IV}_9X^{IV}_{10}C$ (SEQ ID NO: 5)
- 10 wherein each amino acid is indicated by standard one-letter abbreviation, and wherein X^{IV}_1 is E, G, P, N, R, T, W, S, L, H, A, Q or Y; X^{IV}_2 is S, T, E, A, D, G, W, P, L, N, V, Y, R or M; X^{IV}_3 is R, Y, V, Q, E, T, L, P, S, K, M, A or W; X^{IV}_4 is L, M, G, F, W, R, S, V, P, A, D, C or T; X^{IV}_5 is V, T, A, R, S, L, W, C, I, E, P, H, F, D or Q; X^{IV}_6 is E, Y, G, T, Q, M, S, N, A or P; X^{IV}_7 is C, V, D, G, L, W, E, V, I, S, M or A; X^{IV}_8 is S, Y, A, W, P, V, L, Q, G, K, F, I, E or D; X^{IV}_9 is R, W, M, D, H, V, G, A, Q, L, S, E or Y; X^{IV}_{10} is M, L, I, S, V, P, W, F, T, Y, R, or Q.
- 15

96. The method of claim 95, wherein the G-CSF modulator is an agonist for the G-CSFR.

20

97. The method of claim 96, wherein the patient suffers from a depressed neutrophil count.

98. The method of claim 97, wherein the depressed neutrophil count is caused by
- 25 a condition selected from the group consisting of chemotherapy-induced neutropenia, AIDS-induced neutropenia and community-acquired pneumonia-induced neutropenia.

99. The method of claim 95, wherein the G-CSF modulator is an antagonist for the G-CSFR.

100. The method of claim 99, wherein the G-CSF modulator is
5 $\text{NH}_3^+ \text{-LLDICE LKLQECARRCN-COO}^-$ (SEQ ID NO: 208)
 $\text{NH}_3^+ \text{-LLDICE LKLQECARRCN-COO}^-$ (SEQ ID NO: 208).

101. A compound comprising a peptide chain approximately 9 to 40 amino acids
10 in length that binds to G-CSFR and contains a sequence of amino acids of formula (VI)

(VI) $\text{X}^{\text{V}}_1 \text{X}^{\text{V}}_2 \text{X}^{\text{V}}_3 \text{X}^{\text{V}}_4 \text{X}^{\text{V}}_5 \text{X}^{\text{V}}_6 \text{CX}^{\text{V}}_7 \text{X}^{\text{V}}_8$ (SEQ ID NO: 6)

wherein each amino acid is indicated by standard one-letter abbreviation, and wherein
 X^{V}_1 is E, C, Q, V, or Y; X^{V}_2 is E, A, L, M, S, W, or Q; X^{V}_3 is K, R or T; X^{V}_4 is L, A, or V;
 X^{V}_5 is R, A, M, H, E, V, L, G, D, Q, or S; X^{V}_6 is E or V; X^{V}_7 is A or G; X^{V}_8 is R, H, G or

15 L.

102. The compound of claim 101, wherein X^{V}_1 is E.

103. The compound of claim 101, wherein X^{V}_2 is A or L.

20

104. The compound of claim 101, wherein X^{V}_3 is K or R.

105. The compound of claim 101, wherein X^{V}_4 is L.

25

106. The compound of claim 101, wherein X^{V}_6 is E.

107. The compound of claim 101, wherein X^{V}_7 is A.

108. The compound of claim 101, wherein X^{V}_8 is R.

109. The compound of claim 101, wherein the sequence of amino acids is selected from the group consisting of:

- EEKLRECAR (SEQ ID NO: 214);
- EARLAECAR (SEQ ID NO: 215);
- 5 CMKLMECAR (SEQ ID NO: 216);
- ELRLRECAH (SEQ ID NO: 217);
- EAKLHECAR (SEQ ID NO: 218);
- ELKLAECAR (SEQ ID NO: 219);
- EARLEECAR (SEQ ID NO: 220);
- 10 EAKLRECAR (SEQ ID NO: 221);
- ELRLAECAR (SEQ ID NO: 222);
- ESRLAECAR (SEQ ID NO: 223);
- EAKLVECAR (SEQ ID NO: 224);
- ESRLRECAR (SEQ ID NO: 225);
- 15 EAKLAECAR (SEQ ID NO: 226);
- QWRLEECAR (SEQ ID NO: 227);
- QLRLEECAR (SEQ ID NO: 228);
- ELRLEECAR (SEQ ID NO: 229);
- EAKLLECAR (SEQ ID NO: 230);
- 20 EARAGVCAG (SEQ ID NO: 231);
- EAKAGVCAG (SEQ ID NO: 232);
- VARLEECAR (SEQ ID NO: 233);
- ELKLDECAR (SEQ ID NO: 234);
- EWRLQECAR (SEQ ID NO: 235);
- 25 EAKLSECAR (SEQ ID NO: 236);
- EARLSECAR (SEQ ID NO: 237);
- ELKLLECAR (SEQ ID NO: 238);
- ELRLQECGR (SEQ ID NO: 239);

EQKLAECAR (SEQ ID NO: 240);
ELRLQECAR (SEQ ID NO: 241);
ELKLEECAR (SEQ ID NO: 242);
ESRLEECAR (SEQ ID NO: 243);
5 EATVQECAR (SEQ ID NO: 244);
ELKLQECAR (SEQ ID NO: 245);
YSRLEECGR (SEQ ID NO: 246);
ELRLRECAL (SEQ ID NO: 247);
EARLLECAR (SEQ ID NO: 248);
10 ESRLLECAR (SEQ ID NO: 249);
VLKLEECAR (SEQ ID NO: 250);
ESKLAECAR (SEQ ID NO: 251);
ESKLRECAR (SEQ ID NO: 252);
EYKLGECAR (SEQ ID NO: 253);
15 ESRLQECAR (SEQ ID NO: 254);
QARLAECAR (SEQ ID NO: 255);
ELKKQECAR (SEQ ID NO: 256);
ESRLSECAR (SEQ ID NO: 257);
EARLEECGR (SEQ ID NO: 258);
20 ESRLAECGR (SEQ ID NO: 259);
EWRLEECAR (SEQ ID NO: 260);
EARLSECGR (SEQ ID NO: 261);
AARLAECAR (SEQ ID NO: 262);
EWKLAECAR (SEQ ID NO: 263);
25 ESKLEECAR (SEQ ID NO: 264);
DVKLAECAR (SEQ ID NO: 265);
ELQLEECAR (SEQ ID NO: 266); and
EYKLASCAR (SEQ ID NO: 267).

110. The compound of claim 109, wherein the sequence of amino acids is selected from the group consisting of:

- RLSICEEKLRECARGC (SEQ ID NO: 268);
PLTTCEARLAECARQL (SEQ ID NO: 269);
5 LALCMKLMECARRY (SEQ ID NO: 270);
ELVMCELRLRECAHRA (SEQ ID NO: 271);
PLARCEAKLHECARQL (SEQ ID NO: 272);
LLSVCELKLAECARSK (SEQ ID NO: 273);
RLEWCEARLEECARRC (SEQ ID NO: 274);
10 RLRVVEAKLRECARGR (SEQ ID NO: 275);
CVAHLELRLAECARQI (SEQ ID NO: 276);
HLARCESRLAECARQL (SEQ ID NO: 277);
RLALLEAKLVECARRL (SEQ ID NO: 278);
DLFSLESRLRECARRV (SEQ ID NO: 279);
15 AVPVLEAKLAECARRF (SEQ ID NO: 280);
YLQQLQWRLEECARGM (SEQ ID NO: 281);
YLELCQLRLEECARQFN (SEQ ID NO: 282);
ELHICELRLEECARGR (SEQ ID NO: 283);
RVARCELRLAECARKS (SEQ ID NO: 284);
20 YLEVLESRLAECARWK (SEQ ID NO: 285);
EAKLLECARAR (SEQ ID NO: 286);
ELSLCEARAGVCAGSVTK (SEQ ID NO: 287);
ELSLCEAKAGVCAGSVTK (SEQ ID NO: 288);
ALWQCVARLEECARSR (SEQ ID NO: 289);
25 CLKSCELKLDECARRM (SEQ ID NO: 290);
ALQTCEWRLQECARSR (SEQ ID NO: 291);
YISQCEAKLAECARLY (SEQ ID NO: 292);
ELSSCEAKLSECARRW (SEQ ID NO: 293);

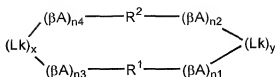
ELSSCEARLSECARRW (SEQ ID NO: 294);
QLLQCELKLEECARQG (SEQ ID NO: 295);
ELLRCEARLAECARGC (SEQ ID NO: 296);
QLRQCELRLQECGRHGN (SEQ ID NO: 297);
5 PLTSCEQKLAECARRF (SEQ ID NO: 298);
LLGMCELRLQECARAK (SEQ ID NO: 299);
ELSRCELKLEECARGM (SEQ ID NO: 300);
DCRPCESRLEECARRL (SEQ ID NO: 301);
RLSVCEARLEECARQL (SEQ ID NO: 302);
10 PLKMCEATVQECARLI (SEQ ID NO: 303);
LLLFCEARLSECARHV (SEQ ID NO: 304);
SLSMCEARLAECARLL (SEQ ID NO: 305);
PLFSCSELKLQECARRCN (SEQ ID NO: 306);
SLERCYSRLEECGRRI (SEQ ID NO: 307);
15 PLTSCSELRLRECALRSN (SEQ ID NO: 308);
KLAACELKLAECARRW (SEQ ID NO: 309);
KLAACELRLAECARRW (SEQ ID NO: 310);
ALTRCELRLAECARKI (SEQ ID NO: 311);
LLQQCELKLAECARSI (SEQ ID NO: 312);
20 QLWQCEARLLEECARRS (SEQ ID NO: 313);
RLRLCESRLLEECARSL (SEQ ID NO: 314);
QLETCVLKLEECARRCN (SEQ ID NO: 315);
ALSQCELRLAECARSVTK (SEQ ID NO: 316);
ELKLAECARRS (SEQ ID NO: 317);
25 ALSRCESKLAECARRQ (SEQ ID NO: 318);
LMSTCESKLRECARSL (SEQ ID NO: 319);
SLQRCEYKLGECARSL (SEQ ID NO: 320);
RLELLESRLQECARQLN (SEQ ID NO: 321);

QMEWCQARLAECARCCN (SEQ ID NO: 322);
PLFSCSELKKQECARRCN (SEQ ID NO: 323);
LLDKCESRLSECARRL (SEQ ID NO: 324);
LLARCEARLEECGRQC (SEQ ID NO: 325);
5 DLLYCESRLAECGRM (SEQ ID NO: 326);
ALQMCEWRLEECARRL (SEQ ID NO: 327);
LLTMCEARLSECGRRL (SEQ ID NO: 328);
ALWRCESRLAECARRS (SEQ ID NO: 329);
LLATCAARLAECARQL (SEQ ID NO: 330);
10 LQTCEWKLAECARSN (SEQ ID NO: 331);
PLRSCSKLEECARQL (SEQ ID NO: 332);
CLRALDVKLAECARHL (SEQ ID NO: 333);
RLKTLELQLEECARRS (SEQ ID NO: 334);
KLRDVELKLAECARRS (SEQ ID NO: 335);
15 SLQRCEYKLASCARSL (SEQ ID NO: 336);
RLARCELRLAECARKS (SEQ ID NO: 337);
DLWYLESKLEECARRCN (SEQ ID NO: 338);
DLWYLESKLEECARRANG (SEQ ID NO: 339);
DLWYLESKLEECARRCNG (SEQ ID NO: 340);
20 KQRELELKLAECAARRS (SEQ ID NO: 341);
QMQEWCARLAECARCCN (SEQ ID NO: 342); and
LLDICEKLQECARRAN (SEQ ID NO: 343).

111. The compound of claim 110, wherein the sequence is:
25 LLDICEKLQECARRAN (SEQ ID NO: 343).

112. The compound of claim 101, comprising a dimer having the structure of
formula (VIII)

(VIII)



- 5 wherein R¹ and R² are independently selected from the sequences of amino acids of formula (V); βA is a β-alanine residue; n1, n2, n3, n4, x and y are independently zero or one with the proviso that the sum of x and y is either one or two; and Lk is a terminal linking moiety selected from the group consisting of a disulfide bond, a carbonyl moiety, a C₁₋₁₂ linking moiety optionally terminated with one or two -NH- linkages and optionally substituted at one or more available carbon atoms with a lower alkyl substituent, a lysine residue or a lysine amide.

113. The compound of claim 101, containing a disulfide bond.

15 114. The compound of claim 113, selected from the group consisting of:

[H]-DLWYLESKLEECARRANG-[NH₂] (SEQ ID NO: 339)

[H]-DLWYLESKLEECARRANG-[NH₂] (SEQ ID NO: 339);

20

[H]-DLWYLESKLEECARRCNG-[NH₂] (SEQ ID NO: 340); and

25

[H]-LLDICEKLQECARRAN-[OH] (SEQ ID NO: 343).

115. The compound of claim 101, wherein the N-terminus of the peptide is coupled to a polyethylene glycol molecule.

116. The compound of claim 101, wherein the N-terminus of the peptide is acetylated.

117. The compound of claim 101, wherein the C-terminus of the peptide is
5 amidated.

118. A pharmaceutical composition comprising a therapeutically effective amount of the compound of claim 101 in combination with a pharmaceutically acceptable carrier.

10

119. A method for treating a patient who would benefit from administration of a G-CSF modulator, comprising administering to the patient a therapeutically effective amount of a compound comprising a peptide chain approximately 9 to 40 amino acids in length that binds to G-CSFR and contains a sequence of amino acids of formula (VI)

15 (VI) $X^V_1X^V_2X^V_3X^V_4X^V_5X^V_6CX^V_7X^V_8$ (SEQ ID NO: 6)

wherein each amino acid is indicated by standard one-letter abbreviation, and wherein X^V_1 is E, C, Q, V, or Y; X^V_2 is E, A, L, M, S, W, or Q; X^V_3 is K, R or T; X^V_4 is L, A, or V; X^V_5 is R, A, M, H, E, V, L, G, D, Q, or S; X^V_6 is E or V; X^V_7 is A or G; X^V_8 is R, H, G or L.

20

120. The method of claim 119, wherein the G-CSF modulator is an agonist for the G-CSFR.

121. The method of claim 120, wherein the patient suffers from a depressed
25 neutrophil count.

122. The method of claim 121, wherein the depressed neutrophil count is caused
a condition selected from the group consisting of chemotherapy-induced neutropenia,
AIDS-induced neutropenia and community-acquired pneumonia-induced neutropenia.

5 123. The method of claim 119, wherein the G-CSF modulator is an antagonist for
the G-CSFR.

124. A compound comprising a peptide chain approximately 10 to 40 amino
acids in length that binds to G-CSFR and contains a sequence of amino acids of

10 formula (VII)

(VII) $X^{VI_1}X^{VI_2}X^{VI_3}X^{VI_4}X^{VI_5}EX^{VI_6}X^{VI_7}X^{VI_8}X^{VI_9}$ (SEQ ID NO: 7)

wherein each amino acid is indicated by standard one-letter abbreviation, and wherein
 X^{VI_1} is A, E or G; X^{VI_2} is E, H or D; X^{VI_3} is R or G; X^{VI_4} is K, Y, M, N, Q, R, D, I, S or E;
 X^{VI_5} is A, S or P; X^{VI_6} is E, D, T, Q, K or A; X^{VI_7} is R, W, K, L, S, A or Q; X^{VI_8} is R or E;
15 and X^{VI_9} is W, G, or R.

125. The compound of claim 124, wherein X^{VI_1} is A.

126. The compound of claim 124, wherein X^{VI_2} is E.

20

127. The compound of claim 124, wherein X^{VI_3} is R.

128. The compound of claim 124, wherein X^{VI_5} is A.

25

129. The compound of claim 124, wherein X^{VI_6} is E.

130. The compound of claim 124, wherein X^{VI_7} is R.

131. The compound of claim 124, wherein X^{VI}_8 is R.

132. The compound of claim 124, wherein and X^{VI}_9 is W.

5 133. The compound of claim 124, wherein the sequence of amino acids is
selected from the group consisting of:

AERKAEERRW (SEQ ID NO: 344);

AERYAEEREG (SEQ ID NO: 345);

AERMAEERRW (SEQ ID NO: 346);

10 AERKAEERRR (SEQ ID NO: 347);

AHRNAEERRW (SEQ ID NO: 348);

AERKSEDWRW (SEQ ID NO: 349);

AERKAEKRR (SEQ ID NO: 350);

AERQAETRRW (SEQ ID NO: 351);

15 AERNAEERRW (SEQ ID NO: 352);

AERQAEERRW (SEQ ID NO: 353);

AERRAEERRW (SEQ ID NO: 354);

AERDAEQRRW (SEQ ID NO: 355);

AERIAEERRW (SEQ ID NO: 356);

20 AERSAEERRW (SEQ ID NO: 357);

AERKAEELRW (SEQ ID NO: 358);

AERKAEESRW (SEQ ID NO: 359);

EERKAEERRW (SEQ ID NO: 360);

ADGKAEERRW (SEQ ID NO: 361);

25 ADGKAEELRW (SEQ ID NO: 362);

ADGMPEERRW (SEQ ID NO: 363);

ADGEAEKRRW (SEQ ID NO: 364);

ADGNAEERRW (SEQ ID NO: 365);

ADGEAEKARW (SEQ ID NO: 366);
AEGEAEKARW (SEQ ID NO: 367);
GERKAEERRW (SEQ ID NO: 368);
AERAEERRW (SEQ ID NO: 369);
5 ADGEAEARRW (SEQ ID NO: 370);
ADGRAEEARW (SEQ ID NO: 371);
AEGRAEEARW (SEQ ID NO: 372);
AERAEKARW (SEQ ID NO: 373);
AERKAEQRW (SEQ ID NO: 374);
10 AERDAEKRRW (SEQ ID NO: 375); and
AERAEKLRW (SEQ ID NO: 376).

134. The compound of claim 133, wherein the sequence of amino acids is selected from the group consisting of:

15 MLAERKAEERRWFNTHGRE (SEQ ID NO: 377);
MLAERKAEERRWFNTHGREK (SEQ ID NO: 378);
GGGMLAERKAEERRWFNTHGRE (SEQ ID NO: 379);
CMLAERKAEERRWFNTHGRE (SEQ ID NO: 380);
CMLAERKAEERRWFNTHGREK (SEQ ID NO: 381);
20 MLAERYAEEREGFNMQWRE (SEQ ID NO: 382);
MLAERMAEERRWFRMG (SEQ ID NO: 383);
IVAERKAEERRRLNTEGHE (SEQ ID NO: 384);
ILAHRNAEERRWFQKHGR (SEQ ID NO: 385);
MLAERKSEDWRWLKTHGRD (SEQ ID NO: 386);
25 MLAERKAEERRLKTQGRE (SEQ ID NO: 387);
ILAERQAETRRWMRNAGSVTK (SEQ ID NO: 388);
MLAERNAEERRWLKRQCG (SEQ ID NO: 389);
MLAERQAEERRWLKMHGGE (SEQ ID NO: 390);

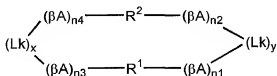
MLAERRAEERRWLKTQGGD (SEQ ID NO: 391);
MLAERQAEERRWLKTQGRD (SEQ ID NO: 392);
MLAERKAEERRWFKTHGRE (SEQ ID NO: 393);
MLAERKAEERRWFNNQGRE (SEQ ID NO: 394);
5 MPAERDAEQRRWLKTHGRE (SEQ ID NO: 395);
ILAERIAEERRWLKTQGR (SEQ ID NO: 396);
MLAERKAEERRWLQTHGRE (SEQ ID NO: 397);
ILAERSAEERRWLKTQGRE (SEQ ID NO: 398);
LLAERKAEELRWLKTHGRE (SEQ ID NO: 399);
10 MLAERKAEERRWLQTHGRE (SEQ ID NO: 400);
MLAERNAEERRW (SEQ ID NO: 401);
MFAERKAEESRWLQSQGRE (SEQ ID NO: 402);
MLEERKAEERRWLKTHGR (SEQ ID NO: 403);
MLAERKAEERRWLKMQGRE (SEQ ID NO: 404);
15 MLAERNAEERRWIFYTHGRE (SEQ ID NO: 405);
MLADGKAEERRWLKTHGLD (SEQ ID NO: 406);
MIADGKAEERRWLKTHGRD (SEQ ID NO: 407);
MLADGKAEELRWLKTQGSD (SEQ ID NO: 408);
MLAERNAEERRWLKTHGRD (SEQ ID NO: 409);
20 MLADGKAEELRWLKTQGRE (SEQ ID NO: 410);
ILADGKAEERRWLKTHGRD (SEQ ID NO: 411);
MLADGMPEERRWLQTHGRD (SEQ ID NO: 412);
MLADGEAEKRRWLNTHGRD (SEQ ID NO: 413);
MLADGNAEERRWLMTHGRD (SEQ ID NO: 414);
25 MLADGEAEKARWLKTQGRE (SEQ ID NO: 415);
MLAEGEAEKARWLKTQGRE (SEQ ID NO: 416);
MLADGKAEERRWLKTQGRE (SEQ ID NO: 417);
MLAERKAEERRWLSAHVRE (SEQ ID NO: 418);

LLGERKAEERRWYKTHARE (SEQ ID NO: 419);
 MLAERKAEERRWLMTHGHD (SEQ ID NO: 420);
 MLAERKAEERRWLKSQCLE (SEQ ID NO: 421);
 LLAEREAEEERRWFKTHGRE (SEQ ID NO: 422);
 5 MLADGEAEARRWFNMHGRE (SEQ ID NO: 423);
 MLADGRAEEARWLKTQGSE (SEQ ID NO: 424);
 MLAEGRAEEARWLKTQGSE (SEQ ID NO: 425);
 MLAERAEKARWLKTQGRE (SEQ ID NO: 426);
 MMAERKAEQRWFDIHGRD (SEQ ID NO: 427);
 10 LTAERDAEKRRWLLTHGGE (SEQ ID NO: 428);
 MLAERQAEERRWLKSQRGE (SEQ ID NO: 429);
 LLAERKAEERRWFATHGRD (SEQ ID NO: 430);
 MLAERAEKLRWLKSQERA (SEQ ID NO: 431);
 MLAERKAEERRWLKTHGGE (SEQ ID NO: 432);
 15 KGGGMLAERKAEERRWFNTHGRE (SEQ ID NO: 490); and
 KSTGGLTAERDAEKRRWLLTHGGE (SEQ ID NO: 491).

135. The compound of claim 124, comprising a dimer having the structure of formula (VIII)

20

(VIII)



wherein R^1 and R^2 are independently selected from the sequences of amino acids of formula (VI); βA is a β -alanine residue; $n1$, $n2$, $n3$, $n4$, x and y are independently zero or one with the proviso that the sum of x and y is either one or two; and Lk is a terminal linking moiety selected from the group consisting of a disulfide bond, a carbonyl moiety, a C_{1-12} linking moiety optionally terminated with one or two $-NH-$ linkages and optionally

substituted at one or more available carbon atoms with a lower alkyl substituent, a lysine residue or a lysine amide.

136. The compound of claim 135, wherein the dimer is selected from the group
5 consisting of:

MLAERKAEERRWFNTHGRE (SEQ ID NO: 377)

MLAERKAEERRWFNTHGRE-K(NH₂) (SEQ ID NO: 378) and

- 10 CMLAERKAEERRWFNTHGRE (SEQ ID NO: 380)

CMLAERKAEERRWFNTHGRE-K (SEQ ID NO: 381).

137. The compound of claim 124, containing a disulfide bond.
15

138. The compound of claim 124, wherein the N-terminus of the peptide is coupled to a polyethylene glycol molecule.

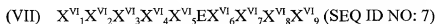
139. The compound of claim 124, wherein the N-terminus of the peptide is
20 acetylated.

140. The compound of claim 124, wherein the C-terminus of the peptide is amidated.

- 25 141. A pharmaceutical composition comprising a therapeutically effective amount of the compound of claim 124 in combination with a pharmaceutically acceptable carrier.

142. A method for treating a patient who would benefit from administration of a
30 G-CSF modulator, comprising administering to the patient a therapeutically effective

amount of a compound comprising a peptide chain approximately 10 to 40 amino acids in length that binds to G-CSFR and contains a sequence of amino acids of formula (VII)



wherein each amino acid is indicated by standard one-letter abbreviation, and wherein

- 5 X^{VI}_1 is A, E or G; X^{VI}_2 is E, H or D; X^{VI}_3 is R or G; X^{VI}_4 is K, Y, M, N, Q, R, D, I, S or E; X^{VI}_5 is A, S or P; X^{VI}_6 is E, D, T, Q, K or A; X^{VI}_7 is R, W, K, L, S, A or Q; X^{VI}_8 is R or E; and X^{VI}_9 is W, G, or R.

- 10 143. The method of claim 142, wherein the G-CSF modulator is an agonist for the G-CSFR.

144. The method of claim 143, wherein the patient suffers from a depressed neutrophil count.

- 15 145. The method of claim 144, wherein the depressed neutrophil count is caused a condition selected from the group consisting of chemotherapy-induced neutropenia, AIDS-induced neutropenia and community-acquired pneumonia-induced neutropenia.

- 20 146. The method of claim 142, wherein the G-CSF modulator is an antagonist for the G-CSFR.

147. A compound comprising a peptide chain approximately 6 to 40 amino acids in length that binds to G-CSF and contains a sequence of amino acids selected from the group consisting of:

- 25 CTWTDLESVY (SEQ ID NO: 433);
HTTNEQFFMC (SEQ ID NO: 434);
DTWLELESRY (SEQ ID NO: 435);
HNSSPMVGVT (SEQ ID NO: 436);

DWQKTIPAYW (SEQ ID NO: 437);
RWGREGLVAALL (SEQ ID NO: 438);
WSGTRVWRCVVT (SEQ ID NO: 439);
MSLLSYLRS (SEQ ID NO: 440);
5 LDLLAI (SEQ ID NO: 441);
RIYGVK (SEQ ID NO: 442);
MIWHMFMSLLF (SEQ ID NO: 443);
FFWASWMHLLW (SEQ ID NO: 444);
FDDCWREREQFLFQAL (SEQ ID NO: 445);
10 CGRASECFRLLM (SEQ ID NO: 446);
RECFQMLER (SEQ ID NO: 447);
CSIRWDFVPGYGLC (SEQ ID NO: 448);
WMQCWDSLSLCYDM (SEQ ID NO: 449);
ALLMCESKLAECARAR (SEQ ID NO: 450);
15 LAHCKKRKEECAAG (SEQ ID NO: 451);
SIDGVYLRTSRT (SEQ ID NO: 452);
SIDGVYLRTSRTRY (SEQ ID NO: 453);
VWRLRGSTLRGLRD (SEQ ID NO: 454);
DRGGGTGVGYWWESY (SEQ ID NO: 455);
20 VWGTVGTWLEY (SEQ ID NO: 456);
LMWVSAY (SEQ ID NO: 457);
RASDEYGALVRFCNTL (SEQ ID NO: 458);
NYWCDSNWWCEIA (SEQ ID NO: 459);
LAHCLLRLEECAAG (SEQ ID NO: 460);
25 LALCLARLRECAGG (SEQ ID NO: 461);
CESRLVECSR (SEQ ID NO: 462);
LLDIAELKLQECARRCN (SEQ ID NO: 463);
KLLDIAELKLQECARRCN (SEQ ID NO: 464);

CSTGGGLTAERDAEKRRWLLTHGGE (SEQ ID NO: 465)
LTAERDAEKRRWLLTHGGEGG (SEQ ID NO: 466);
LTAERDAEKRRWLLTHGGEGGK (SEQ ID NO: 467);
LTAERDAEKRRWLLTHGGEGGGG (SEQ ID NO: 468);
5 LTAERDAEKRRWLLTHGGEGGGGK (SEQ ID NO: 469);
ESGWVW (SEQ ID NO: 470);
NSGWVW (SEQ ID NO: 471);
SGWVW (SEQ ID NO: 472);
PLGKCEATCREMARYFN (SEQ ID NO: 473);
10 SLQRCEYKLASVRGLCN (SEQ ID NO: 474)
DLWYLESKLEEAARRCNG (SEQ ID NO: 475);
PYMGTRSRAKLLRQQ (SEQ ID NO: 476);
RNAGERRWFKTQGWY (SEQ ID NO: 477);
MLAERNADRRWFNTHGRD (SEQ ID NO: 478);
15 MMADGRLRNSVGLILWCD (SEQ ID NO: 479);
MLADGRLRNVVG (SEQ ID NO: 480);
LLADVRRRNGVGLLRMRD (SEQ ID NO: 481);
MLADGRLRNFGG (SEQ ID NO: 482);
TYMTYVYWLC (SEQ ID NO: 483); (CORE 158)
20 RFGERWGL (SEQ ID NO: 484);
HWLWWGWNF (SEQ ID NO: 485);
RECFQMLERC (SEQ ID NO: 486);
ILAHRNAKERRWFQKHGR (SEQ ID NO: 487); and
CSTGGGLTAERDAEKRRWLLTHGGEK (SEQ ID NO: 489).

25

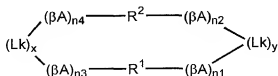
148. The compound of claim 147, wherein the sequence is selected from the group consisting of:

LLDIAELKLEQARRCN (SEQ ID NO: 463); and

149. The compound of claim 147, comprising a dimer having the structure of formula (VIII)

5

(VIII)



wherein R¹ and R² are independently selected from the sequences of amino acids of claim
10 122; β A is a β -alanine residue; n₁, n₂, n₃, n₄, x and y are independently zero or one with
the proviso that the sum of x and y is either one or two; and L_k is a terminal linking
moiety selected from the group consisting of a disulfide bond, a carbonyl moiety, a C₁₋₁₂
linking moiety optionally terminated with one or two -NH- linkages and optionally
substituted at one or more available carbon atoms with a lower alkyl substituent, a lysine
15 residue or a lysine amide.

150. The compound of claim 149, wherein the dimer is selected from the group consisting of:

20

CSTGGGLTAERDAEKRRWLLTHGGE (SEQ ID NO: 465)
|
CSTGGGLTAERDAEKRRWLLTHGGE (SEQ ID NO: 489):

25

LTAERDAEKRRWLLTHGGEGG (SEQ ID NO: 466)
LTAERDAEKRRWLLTHGGEGG-K (SEQ ID NO: 467); and

30

LTAERDAEKRRWLLTHGGEGGGGG (SEQ ID NO: 468)
LTAERDAEKRRWLLTHGGEGGGGG-K (SEQ ID NO: 469).

151. The compound of claim 147, containing a disulfide bond.

152. The compound of claim 151, selected from the group consisting of:

[H]-DLWYLESKLEEAARRCNG-[NH₂] (SEQ ID NO: 475)

[H]-DLWYLESKLEEAARRCNG-[NH₂] (SEQ ID NO: 475);

[H]-LLDIAELKLQECARRCN-[OH] (SEQ ID NO: 463); and

[H]-KLLDIAELKLQECARRCN-[OH] (SEQ ID NO: 464).

153. The compound of claim 147, wherein the N-terminus of the peptide is coupled to a polyethylene glycol molecule.

154. The compound of claim 147, wherein the N-terminus of the peptide is acetylated.

155. The compound of claim 147, wherein the C-terminus of the peptide is amidated.

156. A pharmaceutical composition comprising a therapeutically effective amount of the compound of claim 147 in combination with a pharmaceutically acceptable carrier.

157. A method for treating a patient who would benefit from administration of a G-CSF modulator, comprising administering to the patient a therapeutically effective amount of a compound comprising a peptide chain approximately 6 to 40 amino acids in length that binds to G-CSF and contains a sequence of amino acids selected from the group consisting of:

CTWTDLESVY (SEQ ID NO: 433);

HTTNEQFFMC (SEQ ID NO: 434);

DTWLELESRY (SEQ ID NO: 435);
HNSSPMVGVT (SEQ ID NO: 436);
DWQKTIPAYW (SEQ ID NO: 437);
RWGREGLVAALL (SEQ ID NO: 438);
5 WSGTRVWRCVVT (SEQ ID NO: 439);
MSLLSYLRS (SEQ ID NO: 440);
LDLLAI (SEQ ID NO: 441);
RIYGVK (SEQ ID NO: 442);
MIWHMFMSLLF (SEQ ID NO: 443);
10 FFWASWMHLLW (SEQ ID NO: 444);
FDDCWREREQFLFQAL (SEQ ID NO: 445);
CGRASECFRLLEM (SEQ ID NO: 446);
RECFQMLER (SEQ ID NO: 447);
CSIRWDFVPGYGLC (SEQ ID NO: 448);
15 WMQCWDSLSCYDM (SEQ ID NO: 449);
ALLMCESKLAECARAR (SEQ ID NO: 450);
LAHCKKRKEECAAG (SEQ ID NO: 451);
SIDGVYLRTSRT (SEQ ID NO: 452);
SIDGVYLRTSRTRY (SEQ ID NO: 453);
20 VRWLRGSLRGLRDR (SEQ ID NO: 454);
DRGGGTGVGYWWESY (SEQ ID NO: 455);
VWGTGTWLEY (SEQ ID NO: 456);
LMWVSAY (SEQ ID NO: 457);
RASDEYGALVRFCNTL (SEQ ID NO: 458);
25 NYWCDSNVVCEIA (SEQ ID NO: 459);
LAHCLLRLEECAAG (SEQ ID NO: 460);
LALCLARLRECAGG (SEQ ID NO: 461);
CESRLVECSRM (SEQ ID NO: 462);

LLDIAELKLQECARRCN (SEQ ID NO: 463);
KLLDIAELKLQECARRCN (SEQ ID NO: 464);
CSTGGGLTAERDAEKRRWLLTHGGE (SEQ ID NO: 465);
LTAERDAEKRRWLLTHGGEGG (SEQ ID NO: 466);
5 LTAERDAEKRRWLLTHGGEGGK (SEQ ID NO: 467);
LTAERDAEKRRWLLTHGGEGGGGG (SEQ ID NO: 468);
LTAERDAEKRRWLLTHGGEGGGGGK (SEQ ID NO: 469);
ESGWVW (SEQ ID NO: 470);
NSGWVW (SEQ ID NO: 471);
10 SGWVW (SEQ ID NO: 472);
PLGKCEATCREMARYFN (SEQ ID NO: 473);
SLQRCEYKLASVRGLCN (SEQ ID NO: 474);
DLWYLESKLEEAARRCNG (SEQ ID NO: 475);
PYMGTRSRAKLLRQQ (SEQ ID NO: 476);
15 RNAGERRWFKTQGWY (SEQ ID NO: 477);
MLAERNADDRRWFNTHGRD (SEQ ID NO: 478);
MMADGRLRNSVGLILWCD (SEQ ID NO: 479);
MLADGRLRNVVG (SEQ ID NO: 480);
LLADVRRRNGVGLLRMGRD (SEQ ID NO: 481);
20 MLADGRLRNFGG (SEQ ID NO: 482);
TYMTYVYWLC (SEQ ID NO: 483);
RFGERWGL (SEQ ID NO: 484);
HWLWWGWNF (SEQ ID NO: 485);
RECFQMLERC (SEQ ID NO: 486);
25 ILAHRNAKERRWFQKHGR (SEQ ID NO: 487); and
CSTGGGLTAERDAEKRRWLLTHGGEK (SEQ ID NO: 489).

159. The method of claim 158, wherein the patient suffers from a depressed
5 neutrophil count.

160. The method of claim 159, wherein the depressed neutrophil count is caused by a condition selected from the group consisting of chemotherapy-induced neutropenia, AIDS-induced neutropenia and community-acquired pneumonia-induced neutropenia.

10

161. The method of claim 157, wherein the G-CSF modulator is an antagonist for the G-CSFR.